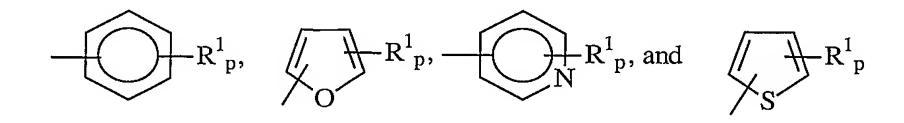
WHAT IS CLAIMED IS:

1. A 3,6-substituted pyran group-containing

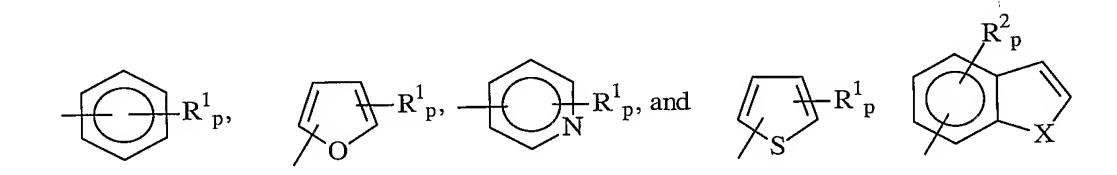
2 compound having the structural formula:

- 3 wherein
- A, A', and B are individually selected from the group of optionally substituted C_{4}
- 5 C₁₄ aryl and heteroaryl wherein heteroatoms of heteroaryl A and/or A' are selected
- from the group consisting of O, N, and S;
- Z is selected from the group consisting of a chemical bond and -Y-($CH_{2)o}$ wherein
- 8 Y is NH or O and o is 0, 1, 2, 3, or 4;
- 9 R is H or C_{1-8} alkyl;
- W is selected from the group consisting of hydrogen and -OH; and
- n and m individually are 0, 1, 2, 3, or 4, and wherein any carbon of $-(CH_2)_n$ may
- be substituted by OR^4 wherein R^4 is C_{1-8} alkyl, C_{2-18} alkylene, or -COOR⁵ wherein
- R⁵ is C_{1-18} alkyl or C_{2-18} alkylene, or a pharmaceutically acceptable derivative or salt
- 14 thereof.
- 1 2. The compound of claim 1, wherein at least one of A and A'
- 2 are selected from the group consisting of:

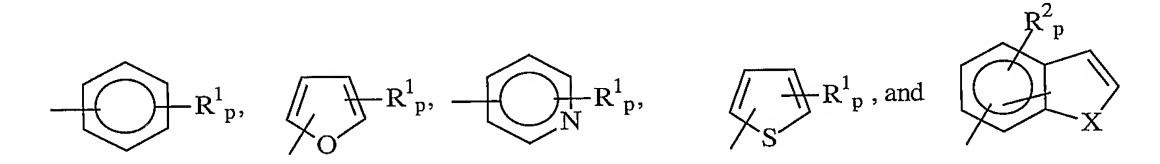


3 where R^1 is selected from the group consisting of C_{1-4} alkyl, C_{2-6} alkenyl, C_{2-6}

- optionally halogenated alkynyl, C₂₋₆ hydroxyalkynyl, halo, -CN, -COOR, where R
- is C_{1-18} alkyl, C_{5-10} cycloalkyl, C_{2-18} alkenyl, -OH, -NO₂, -NH₂, -OR² where R² is
- 6 C_{1-8} alkyl, C_{5-6} cycloalkyl, or C_{2-8} alkenyl.
- 1 3. The compound of claim 1, wherein B is selected from the
- 2 group



- 3 where R^1 is selected from the group consisting of C_{1-4} alkyl, C_{2-6} alkenyl, C_{2-6}
- optionally halogenated alkynyl, C₂₋₆ hydroxyalkynyl, halo, -CN, -COOR, where R
- is C_{1-18} alkyl, C_{5-10} cycloalkyl, C_{2-18} alkenyl, -OH, -NO₂, -NH₂, -OR² where R² is C_{1-8}
- 6 alkyl, C_{5-6} cycloalkyl, or C_{2-8} alkenyl; and
- wherein R² have the meaning of R¹ and also a 5 or 6 membered heterocycle
- 8 containing 1 or more heteroatoms selected from the group consisting of N, O, and
- 9 S, and wherein X is N, O, or S.
- 1 4. The compound of claim 2, wherein B is selected from the
- 2 group



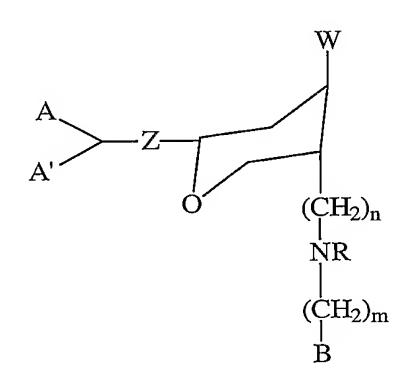
- 3 where R^1 is selected from the group consisting of C_{1-4} alkyl, C_{2-6} alkenyl, C_{2-6}
- optionally halogenated alkynyl, C₂₋₆ hydroxyalkynyl, halo, -CN, -COOR, where R
- is C_{1-18} alkyl, C_{5-10} cycloalkyl, C_{2-18} alkenyl, -OH, -NO₂, -NH₂, -OR² where R² is C_{1-8}
- 6 alkyl, C_{5-6} cycloalkyl, or C_{2-8} alkenyl; and

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wherein R² have the meaning of R¹ and also a 5 or 6 membered heterocycle

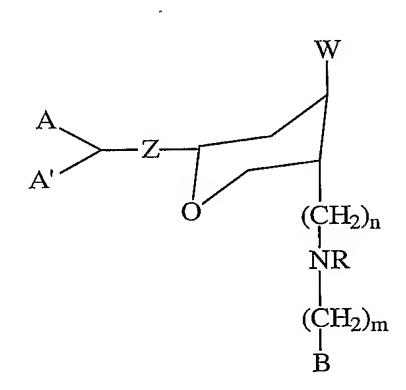
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- 8 containing 1 or more heteroatoms selected from the group consisting of N, O, and
- 9 S, and wherein X is N, O, or S.
- The compound of claim 3, wherein A and A' are both
- 2 unsubstituted phenyl.
- 1 6. The compound of claim 1, having the formula



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7. The compound of claim 2, having the formula

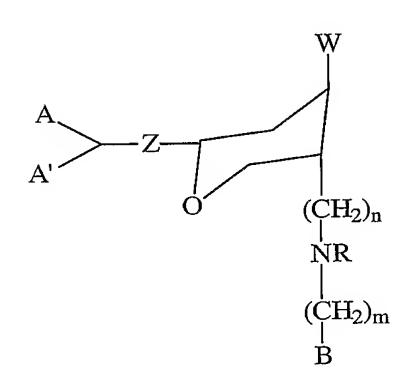


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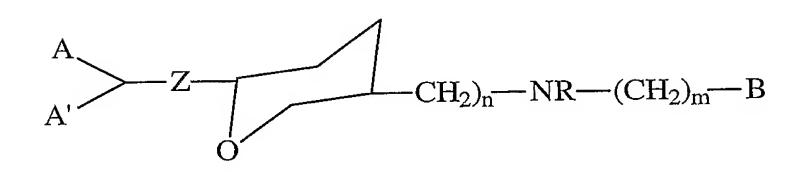
8. The compound of claim 3, having the formula



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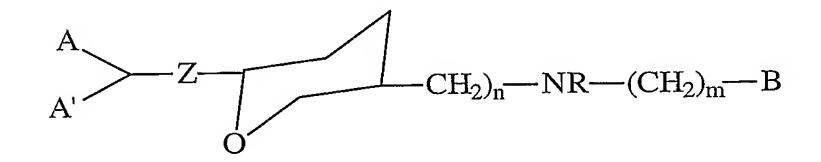
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9. The compound of claim 1, having the formula



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10. The compound of claim 2, having the formula



1 11. The compound of claim 3, having the formula

$$A$$
 Z
 $CH_2)_n$
 NR
 $CH_2)_m$
 B

- 1 12. The compound of claim 1, having a formula selected from the
- 2 group consisting of:

- 1 13. The compound of claim 2, having a formula selected from the
- 2 group consisting of:

1 14. The compound of claim 3, having a formula selected from the

2 group consisting of:

- 1 15. The compound of claim 5, having a formula selected from the
- 2 group consisting of:

- 1 16. The compound of claim 1, selected from the group consisting
- 2 of:
- 3 Synthesis of Cis-(6-benzhydryl-tetrahydropyran-3-yl)-(4-hydroxy-benzyl)-amine (16h);
- 4 Synthesis of Cis-(6-benzhydryl-tetrahydropyran-3-yl)-(1H-iodo-5-ylmethyl)-amine (16n);
- 5 Synthesis of Cis-(6-benzhydryl-tetrahydropyran-3-yl)-(4-amino-benzyl)-amine (160);
- 6 Synthesis of Cis-(6-benzhydryl-tetrahydropyran-3-yl)-(3,4-dichloro-benzyl)-amine (16i);
- Procedure E. Synthesis of (2S, 4R, 5R)-2-benzhydryl-5-(4-methoxy-benzylamino)-
- 8 tetrahydropyran-4-ol (-)29a;
- 9 Synthesis of (2S, 4R, 5R)-2-benzhydryl-5-(4-fluoro-benzylamino)-tetrahydro-pyran-4-ol
- 10 (-)29b;
- Synthesis of (2S, 4R, 5R)-2-benzhydryl-5-benzylamino-tetrahydro-pyran-4-ol (-)29d;

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Synthesis of (2S, 4R, 5R)-2-benzhydryl-5-(2,4-dimethoxy-benzylamino)-tetrahydropyran-4-

14 ol (-)-29e;

15

- Synthesis of (2S, 4R, 5R)-2-benzhydryl-5-(3,5-dimethoxy-benzylamino)-tetrahydropyran-4-
- 17 ol (-)-29f;
- Procedure H. Synthesis of (2S, 4R, 5R)-2-benzhydryl-5-(4-hydroxy-benzylamino)-
- tetrahydropyran-4-ol (-)32a;
- Synthesis of (2S, 4R, 5R)-2-benzhydryl-5-[(1H-indol-5-ylmethyl)-amino]-tetrahydropyran-
- 21 4-ol (-)32b;
- Synthesis of (2R, 4S, 5S)-2-benzhydryl-5-(4-hydroxy-benzylamino)-tetrahydro-pyran-4-ol
- (+)32a;
- Synthesis of (2R, 4S, 5S)-2-benzhydryl-5-[(1H-indol-5-ylmethyl)-amino]-tetrahydropyran-4-
- 25 ol (+)32b;
- Synthesis of cis-(3S, 6S)-(6-benzhydryl-tetrahydropyran-3-yl)-(4-hydroxy-benzyl)-amine
- 27 (-)37a; and
- Synthesis of cis-(3R, 6R)-(6-benzhydryl-tetrahydropyran-3-yl)-(4-hydroxy-benzyl)-
- 29 amine (+)37a.
- 1 The compound of claim 1, selected from the group consisting
- 2 of:
- Procedure E. Synthesis of (2S, 4R, 5R)-2-benzhydryl-5-(4-methoxy-benzylamino)-
- 4 tetrahydropyran-4-ol (-)29a;
- 5 Synthesis of (2S, 4R, 5R)-2-benzhydryl-5-(4-fluoro-benzylamino)-tetrahydro-pyran-4-ol
- 6 (-)29b;
- 7 Synthesis of (2S, 4R, 5R)-2-benzhydryl-5-benzylamino-tetrahydro-pyran-4-ol (-)29d;

8

- 9 Synthesis of (2S, 4R, 5R)-2-benzhydryl-5-(2,4-dimethoxy-benzylamino)-tetrahydropyran-4-
- 10 ol (-)-29e;
- Synthesis of (2S, 4R, 5R)-2-benzhydryl-5-(3,5-dimethoxy-benzylamino)-tetrahydropyran-4-
- 12 ol (-)-29f;

Procedure H. Synthesis of (2S, 4R, 5R)-2-benzhydryl-5-(4-hydroxy-benzylamino)-

- tetrahydropyran-4-ol (-)32a;
- Synthesis of (2S, 4R, 5R)-2-benzhydryl-5-[(1H-indol-5-ylmethyl)-amino]-tetrahydropyran-
- 16 4-ol (-)32b;
- Synthesis of (2R, 4S, 5S)-2-benzhydryl-5-(4-hydroxy-benzylamino)-tetrahydro-pyran-4-ol
- 18 (+)32a;
- Synthesis of (2R, 4S, 5S)-2-benzhydryl-5-[(1H-indol-5-ylmethyl)-amino]-
- 20 tetrahydropyran-4-ol (+)32b;
- Synthesis of *cis*-(3S, 6S)-(6-benzhydryl-tetrahydropyran-3-yl)-(4-hydroxy-<u>benzyl</u>)-amine
- 22 (-)37a; and
- 23 Synthesis of cis-(3R, 6R)-(6-benzhydryl-tetrahydropyran-3-yl)-(4-hydroxy-benzyl)-amine
- (+)37a.
- 1 18. A method of reducing monoamine reuptake in a mammalian
- 2 species, comprising administering a binding amount of a monoamine receptor binder
- 3 comprising a compound of claim 1.
- 1 19. A method of reducing monoamine reuptake in a mammalian
- 2 species, comprising administering a binding amount of a monoamine receptor binder
- 3 comprising a compound of claim 2.
- 1 20. A method of reducing monoamine reuptake in a mammalian
- 2 species, comprising administering a binding amount of a monoamine receptor binder
- 3 comprising a compound of claim 12.
- 1. A method for the treatment of depression, comprising
- 2 administering to a patient exhibiting signs of depression, a compound of claim 1 in
- an amount effective to inhibit reuptake of serotonin at the SERT and norepinephrine
- 4 at the NET.
- 1 22. The method of claim 21 wherein the compound exhibits
- 2 greater inhibition of serotonin and norepinephrine reuptake than of dopamine
- 3 reuptake.

23. A method for the treatment of depression, comprising administering to a patient exhibiting signs of depression, a compound of claim 1 in an amount effective to inhibit norepinephrine reuptake at the NET.

1 24. The method of claim 23 wherein said compound exhibits 2 higher norepinephrine reuptake inhibition than serotonin reuptake inhibition and 3 dopamine reuptake inhibition.